

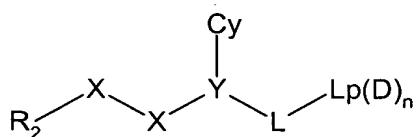
Continuation of Serial No. 10/030,187

Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of Claims:

1 (currently amended) : A serine protease inhibitor compound of formula (I)



(I)

wherein:

R_2 is:-

(i) phenyl optionally being substituted in the 3 and/or 4 position by halo, nitro, thiol, haloalkoxy, hydrazido, alkylhydrazido, amino, cyano, haloalkyl, alkylthio, alkenyl, alkynyl, acylamino, tri or difluoromethoxy, carboxy, acyloxy, MeSO_2^- or R_1 , and optionally substituted at the 6 position by amino, hydroxy, halo, alkyl, carboxy, alkoxy carbonyl, cyano, amido, aminoalkyl, alkoxy or alkylthio;

(ii) naphth-2-yl optionally substituted at the 6 or 7 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R_{1j} and optionally substituted at the 3 position by amino, hydroxy, halo, alkyl, carboxy, cyano, amido, aminoalkyl, alkoxy or alkylthio;

(iii) isoquinolin-7-yl, indol-5-yl, indol-6-yl, indazol-5-yl, indazol-6-yl, benzothiazol-6-yl or benzisoxazol-5-yl optionally substituted at the 3 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R_{1j} ;

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- (iv) benzimidazol-5-yl or benzothiazol-6-yl optionally substituted at the 2 position by amino;
- (v) thien-2-yl or thien-3-yl optionally substituted at the 4 or 5 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R₁;
- (vi) 3,4-methylenedioxyphenyl, 2,3-dihydroindol-6-yl, 3,3-dichloro-2-oxo-indol-6-yl or 1-methyl-3-aminoindazol-5-yl;
- (vii) benzothiazol-2-yl, imidazo[1,2-a]pyrimidin-2-yl or tetrahydroimidazo[1,2-a]pyrimidin-2-yl;
- (viii) pyrazol-2-yl optionally substituted at the 5 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R₁;
- (ix) pyrid-2-yl optionally substituted at the 5 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R₁;
- (x) pyrid-3-yl optionally substituted at the 6 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R₁;
- (xi) benzofur-2-yl optionally substituted at the 3 position by amino, hydroxy, halo, alkyl, carboxy, cyano, amido, aminoalkyl, alkoxy or alkylthio and at the 5 or 6 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R_{1j};
- (xii) indol-2-yl optionally substituted on the indole nitrogen atom by alkyl and optionally substituted at the 5 or 6 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R_{1j};
- (xiii) indol-6-yl substituted at the 5 position by amino, hydroxy, halo, alkyl, carboxy, alkoxy carbonyl, cyano, amido, aminoalkyl, alkoxy or alkylthio and optionally substituted at the 3 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R_{1j}; or

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(xiv) benzo[b]thiophen-2-yl optionally substituted at the 3 position by amino, hydroxy, halo, alkyl, carboxy, cyano, amido, aminoalkyl, alkoxy or alkylthio and at the 5 or 6 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R_{1j}; a 5 or 6 membered aromatic carbon ring optionally interrupted by a nitrogen, oxygen or sulphur ring atom, optionally being substituted in the 3 and/or 4 position (in relation to the point of attachment of X-X) by halo, nitro, thiol, haloalkoxy, hydrazido, alkylhydrazido, amino, cyano, haloalkyl, alkylthio, alkenyl, alkynyl, acylamino, tri- or difluoromethoxy, carboxy, acyloxy, MeSO₂- or R₁, or the substituents at the 3 and 4 positions taken together form a fused ring which is a 5 or 6 membered carbocyclic or heterocyclic ring optionally substituted by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R_{1j}, and optionally substituted in the position alpha to the X-X group (i.e. 6 position for a six membered aromatic ring etc) by amino, hydroxy, halo, alkyl, carboxy, alkoxycarbonyl, cyano, amido, aminoalkyl, alkoxy or alkylthio

with the proviso that R₂ cannot be aminoisoquinolyl; -X-X- is -CONH- each X independently is a C, N, O or S atom or a CO, CR_{1a}, C(R_{1a})₂ or NR_{1a} group, at least one X being C, CO, CR_{1a} or C(R_{1a})₂;

each R_{1a} independently represents hydrogen or hydroxyl, alkoxy, alkyl, aminoalkyl, hydroxyalkyl alkoxylalkyl, alkoxycarbonyl, alkylaminocarbonyl, alkoxycarbonylamine, acyloxymethoxy carbonyl or alkylamino optionally substituted by hydroxy, alkylamino, alkoxy, exo, aryl or cycloalkyl;

R₁ is hydrogen, hydroxy, alkoxy, alkyl, alkylaminoalkyl, alkanoyl, hydroxyalkyl, alkoxyalkyl, alkoxycarbonyl, alkylaminocarbonyl, alkylamino, carboxyl, carboxymethyl, amido or amidomethyl;

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R_{1j} is hydrogen, hydroxy, alkoxy, alkyl, alkanoyl, hydroxyalkyl, alkoxyalkyl, alkoxycarbonyl, alkylamino, carboxyl, carboxymethyl, amido or amidomethyl; as defined for R_{1a} , provided that R_1 is not unsubstituted aminoalkyl;

Y (the α -atom) is a CH nitrogen atom or a CR_{1b} group;

C_y is an optionally R_{3a} substituted: phenyl, pyridyl, thienyl, thiazolyl, naphthyl, piperidinyl, furanyl, pyrrolyl, isoxazolyl, isothiazolyl, pyrazolyl, oxazolyl, imidazolyl, 1,2,4-thiadiazolyl, 1,3,4-thiadiazolyl, pyrimidinyl, pyridazinyl, quinolyl, isoquinolyl, benzofuryl, benzothienyl or cycloalkyl group, or a phenyl group substituted by $R_{3i}X_i$; a saturated or unsaturated, mono or poly cyclic, homo or heterocyclic group, optionally substituted by groups R_{3a} or $R_{3i}X_i$;

each R_{3a} independently is hydrogen, hydroxyl, alkoxy, aralkyloxy, alkyl, alkylaminoalkyl, hydroxymethyl, carboxy, alkoxyalkyl, alkoxycarbonyl, alkylaminocarbonyl, aminomethyl, $CONH_2$, CH_2CONH_2 , (1-6C) alkanoylamino, alkoxycarbonylamino R_{1e} , amino, halo, cyano, nitro, thiol, alkylthio, alkylsulphonyl, alkylsulphenyl, triazolyl, imidazolyl, tetrazolyl, hydrazido, alkylimidazolyl, thiazolyl, alkylthiazolyl, alkylloxazolyl, oxazolyl, alkylsulphonamido, alkylaminosulphonyl, aminosulphonyl, haloalkoxy, haloalkyl, a group of the formula $-C(X^3)N(R^{11})R^{12}$ (wherein X^3 is O or S; and R^{11} and R^{12} are independently selected from hydrogen, methyl or ethyl or together with the nitrogen atom to which they are attached form a pyrrolidin-1-yl, piperidin-1-yl or morpholino group), or $-OCH_2O-$ which is bonded to two adjacent ring atoms in C_y ;

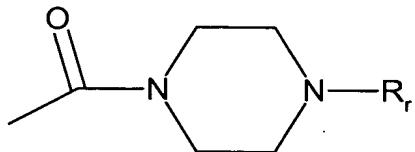
X_i is a bond, O, NH or CH_2 ;

R_{3i} is phenyl, pyridyl or pyrimidinyl optionally substituted by R_{3a} ;

R_{1b} , R_{1e} and R_{1j} are as defined for R_{1a} , and

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-L-Lp(D)_n is of the formula:

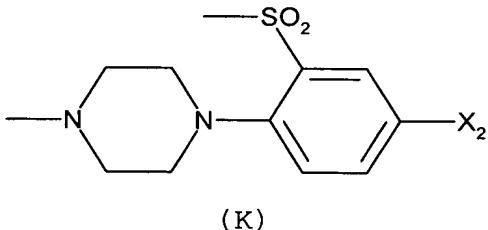


in which R_r is -(CH₂)_c-R_c, -CHReR_f, -CH₂-CHReR_f, -CH₂-CH₂-CHReR_f, or R_g in which c is 1 or 2; R_c is thieryl, thiazolyl (which may bear an amino substituent), isothiazolyl, oxazolyl, isoxazolyl, pyrazolyl, imidazolyl, pyridyl (which may bear an alkylsulphonyl, aminosulphonyl, alkylaminosulphonyl, alkylaminocarbonyl, amino, amido, (1-4C)alkoxycarbonyl, carboxy, acetylarnino, chloro, fluoro, cyano, (1-3C)alkyl, trifluoromethyl, methoxy, ethoxy, nitro, hydroxy, alkylsulphonylamino, triazolyl or tetrazolyl substituent), pyrimidinyl, pyridazinyl, pyrazinyl or phenyl (which may bear a methyl, methylarnino, dimethylarnino, carboxy, dialkylaminosulphonyl, alkylsulphonyl, aminosulphonyl, alkylaminosulphonyl, alkylaminocarbonyl, amino, amido, alkoxy carbonyl, acetylarnino, chloro, fluoro, cyano, methoxy, ethoxy, nitro, hydroxy, alkylsulphonylamino, triazolyl or tetrazolyl substituent); each of R_e and R_f independently is hydrogen or C₁₋₃alkyl; or CHReR_f is cyclopentyl (which may bear a hydroxy, amino, (1-3C)alkoxy, (1-3C)hydroxyalkyl, (1-3C)alkyl, carboxy, methoxycarbonyl or ethoxycarbonyl substituent at the 3- or 4-position), cyclohexyl (which may bear a hydroxy, amino, (1-3C)alkoxy, (1-3C)hydroxyalkyl, (1-3C)alkyl, carboxy, methoxycarbonyl or ethoxycarbonyl substituent at the 3- or 4-position), tetrahydropyran-4-yl, tetrahydrothiopyran-4-yl, pyrrolidin-3-yl (which may bear a hydroxy, amino, (1-3C)alkoxy, (1-3C)hydroxyalkyl, (1-3C)alkyl, carboxy, methoxycarbonyl or ethoxycarbonyl substituent at the 1-position), piperidin-4-yl (which may bear a hydroxy, amino, (1-3C)alkoxy, (1-3C)hydroxyalkyl, (1-3C)alkyl, carboxy, methoxycarbonyl or ethoxycarbonyl substituent at the 1-

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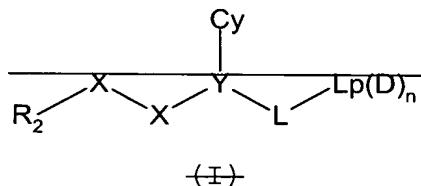
position), or indan-2-yl; and R_g is 2-methylsulphonylphenyl which may bear a 4-fluoro substituent or R_g is 1,1-dioxobenzo[b]thiophen-7-yl;

or a physiologically-tolerable salt thereof;
provided that Lp(D)_n is not of the formula (K):



wherein X₂ is fluoro or hydrogen.

2 (currently amended): A compound according to claim 1
~~serine protease inhibitor compound of formula (I)~~



wherein:

R₂ is a 5 or 6 membered aromatic carbon ring optionally interrupted by a nitrogen, oxygen or sulphur ring atom, optionally being substituted in the 3 and/or 4 position (in relation to the point of attachment of X-X) by halo, nitro, thiol, haloalkoxy, hydrazido, alkylhydrazido, amino, cyano, haloalkyl, alkylthio, alkenyl, alkynyl, acylamino, tri or difluoromethoxy, carboxy, acyloxy, MesO₂ or R₁, or the substituents at the 3 and 4 positions taken together form a fused ring which is a 5 or 6 membered carbocyclic or heterocyclic ring optionally substituted by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R₁, and optionally substituted in the position alpha to the X-X group (i.e. 6 position for a six membered

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aromatic ring etc) by amino, hydroxy, halo, alkyl, carboxy, alkoxy carbonyl, cyano, amide, aminoalkyl, alkoxy or alkylthio with the proviso that R_2 cannot be aminoisoquinolyl;

each X independently is a C, N, O or S atom or a CO, CR_{1a} , $C(R_{1a})_2$ or NR_{1a} group, at least one X being C, CO, CR_{1a} or $C(R_{1a})_2$;

each R_{1a} independently represents hydrogen or hydroxyl, alkoxy, alkyl, aminoalkyl, hydroxyalkyl, alkoxyalkyl, alkoxy carbonyl, alkylaminocarbonyl, alkoxy carbonyl amino, acyloxymethoxy carbonyl or alkylamino optionally substituted by hydroxy, alkylamino, alkoxy, exo, aryl or cycloalkyl;

R_1 is as defined for R_{1a} , provided that R_1 is not unsubstituted aminoalkyl;

Y (the α atom) is a nitrogen atom or a CR_{1b} group;

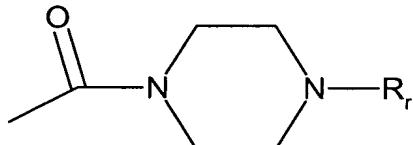
Cy is an optionally R_{3a} substituted: phenyl, pyridyl, thienyl, thiazolyl, naphthyl, piperidinyl or cycloalkyl group; a saturated or unsaturated, mono or poly cyclic, homo or heterocyclic group, optionally substituted by groups R_{3a} or phenyl optionally substituted by R_{3a} ;

each R_{3a} independently is hydrogen, hydroxyl, methoxy, ethoxy, methyl, ethyl, methylaminomethyl, dimethylaminomethyl, hydroxymethyl, carboxy, methoxymethyl, methoxycarbonyl, ethoxycarbonyl, methylaminocarbonyl, dimethylamino-carbonyl, aminomethyl, $CONH_2$, CH_2CONH_2 , acetyl amino, methoxycarbonyl amino, ethoxycarbonyl amino, t-butoxycarbonyl amino, amino, fluoro, chloro, cyano, nitro, thiol, methylthio, methylsulphonyl, ethylsulphonyl, methylsulphenyl, methylsulphonylamido, ethylsulphonylamido, methylaminosulphonyl, ethylaminosulphonyl, aminosulphonyl, trifluoromethoxy or trifluoromethyl; R_1e , amino, halo, cyano, nitro, thiol, alkylthio, alkylsulphonyl, alkylsulphenyl, triazolyl, imidazolyl, tetrazolyl, hydrazido, alkyl imidazolyl, thiazolyl, alkyl thiazolyl, alkyl oxazolyl,

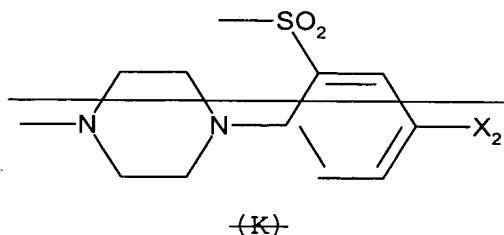
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~~oxazolyl, alkylsulphonamide, alkylaminosulphonyl,~~
~~aminosulphonyl, haloalkoxy and haloalkyl;~~

~~R_{1b}, R_{1e} and R_{1j} are as defined for R_{1a}; and~~
~~-L-Lp(D)_n is of the formula:~~

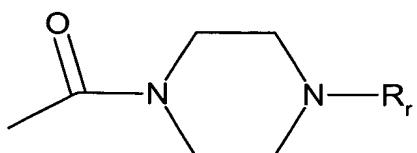


in which R_r is -(CH₂)_c-R_C, -CHR_eR_f, -CH₂-CHR_eR_f, or R_g in
which c is 1 or 2; R_C is pyridyl or phenyl (which phenyl may
bear a fluoro, chloro, methyl, CONH₂, SO₂NH₂,
methylaminosulphonyl, dimethylaminosulphonyl, methoxy or
methylsulphonyl substituent); each of R_e and R_f independently
is hydrogen or C₁₋₃alkyl; or CHR_eR_f is cyclopentyl (which may
bear a methyl, ethyl or hydroxymethyl substituent at the 3- or
4-position), cyclohexyl (which may bear a methyl, ethyl or
hydroxymethyl, (1-3C)alkyl, carboxy, methoxycarbonyl or
ethoxycarbonyl substituent at the 3- or 4-position),
tetrahydropyran-4-yl, tetrahydrothiopyran-4-yl, pyrrolidin-3-
yl (which may bear a 1-methyl substituent), piperidin-4-yl
(which may bear a 1-methyl substituent) or indan-2-yl; and R_g
is 2-methylsulphonylphenyl which may bear a 4-fluoro
substituent or R_g is λ⁶-1,1-dioxobenzo[b]thiophen-7-yl+
~~or a physiologically tolerable salt thereof;~~
~~provided that Lp(D)n is not of the formula (K):~~



wherein X₂ is fluoro or hydrogen.

3 (original): A compound according to claim 1 wherein -L-Lp(D)_n is of the formula:



in which R_r is -(CH₂)_c-R_C; in which c is 2; R_C is thienyl, thiazolyl (which may bear an amino substituent), isothiazolyl, oxazolyl, isoxazolyl, pyrazolyl, imidazolyl, pyridyl (which may bear an amino, methoxycarbonyl, carboxy, fluoro, cyano, methyl, methylsulphonyl, aminosulphonyl, methylaminosulfonyl, dimethylaminosulfonyl, or trifluoromethyl substituent), pyrimidinyl, pyridazinyl, pyrazinyl or phenyl (which phenyl may bear a fluoro, chloro, cyano, methyl, amino, methylsulphonyl, aminosulphonyl, methylaminosulphonyl, dimethylaminosulphonyl, methylamino, dimethylamino, carboxy, methoxycarbonyl or methoxy substituent).

4 (currently amended): A compound according to claim ~~any one of claims 1 to 3~~ wherein R_C is thiazolyl (which may bear an amino substituent), pyrimidinyl, pyrazolyl, imidazolyl, pyridyl (which may bear a methylsulphonyl, aminosulphonyl, methylaminosulfonyl, dimethylaminosulfonyl, fluoro, cyano, methyl or trifluoromethyl substituent), pyridazinyl, pyrazinyl or phenyl (which phenyl may bear a fluoro, chloro, cyano, methyl, amino, methylamino, dimethylamino, carboxy, methoxycarbonyl, methylsulphonyl, aminosulphonyl, methylaminosulfonyl, dimethylaminosulfonyl, or methoxy substituent).

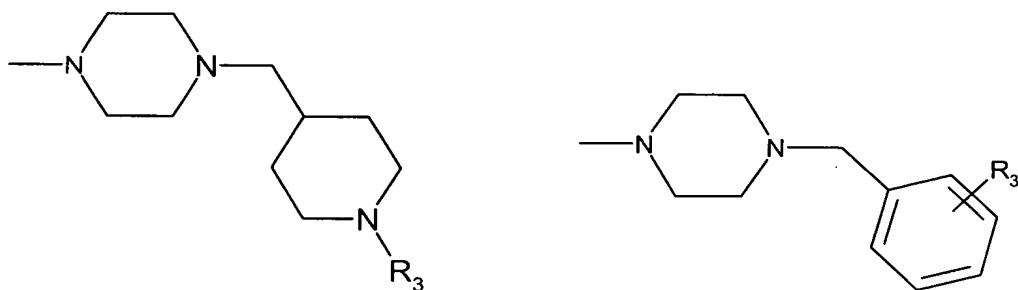
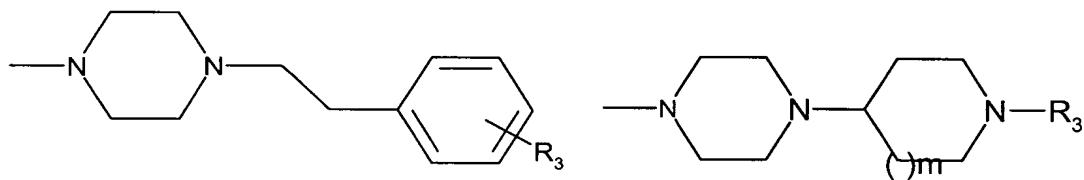
5 (currently amended): A compound according to claim ~~any one of claims 1 to 4~~ wherein R_C is thiazolyl (which may bear an amino substituent), pyrazolyl, imidazolyl, pyridyl (which may

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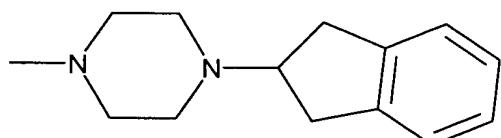
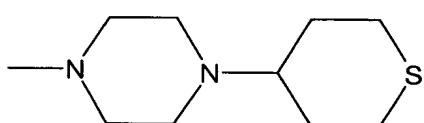
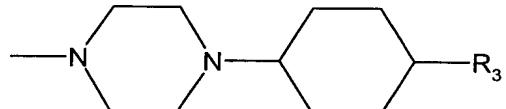
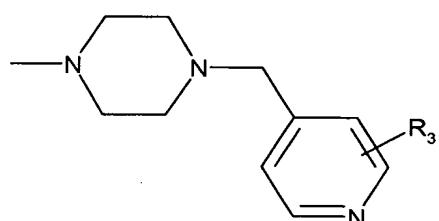
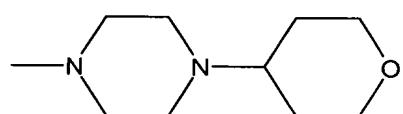
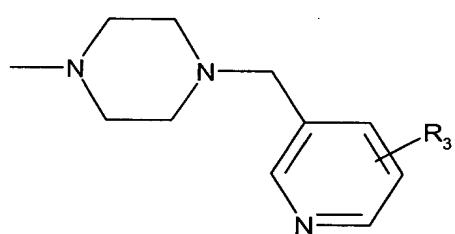
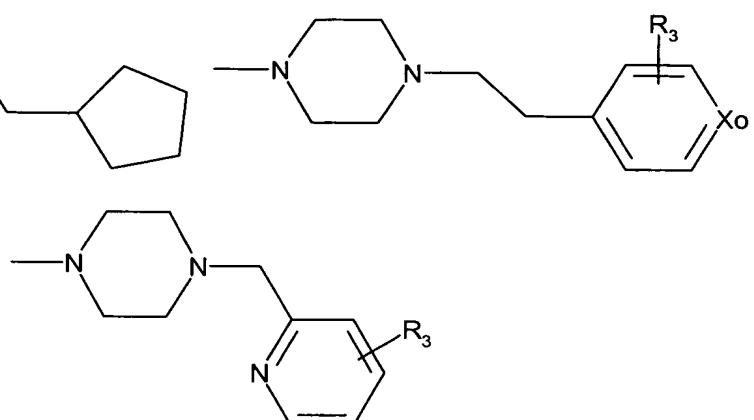
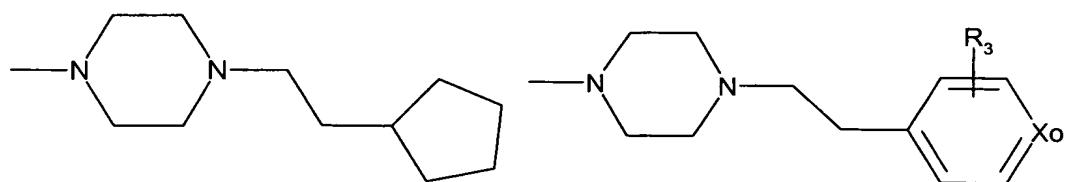
bear a fluoro, cyano, methyl or trifluoromethyl substituent), pyridazinyl or pyrazinyl.

6 (currently amended): A compound according to claim ~~any one~~ of claims 1 to 5 wherein Rc is thiazol-2-yl, 2-aminothiazol-4-yl, pyrazol-1-yl, pyrazol-4-yl, pyridazin-3-yl, imidazol-1-yl, imidazol-4-yl, pyrazin-2-yl, pyrid-2-yl, pyrid-3-yl, pyrid-4-yl, 3-fluoropyrid-4-yl, 2-cyanopyrid-4-yl, 2-methylpyrid-4-yl or 2-trifluoromethylpyrid-6-yl.

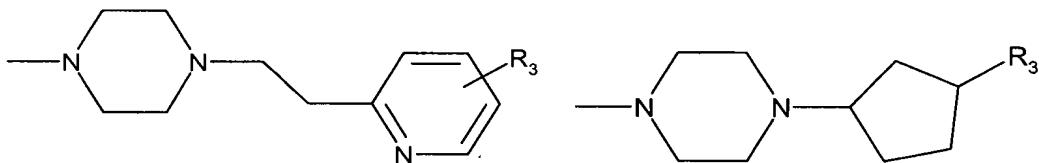
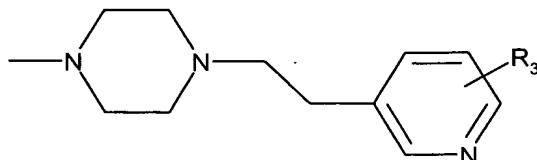
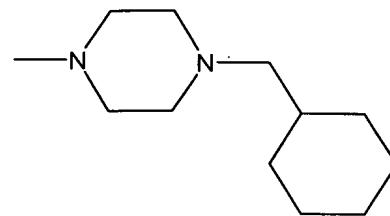
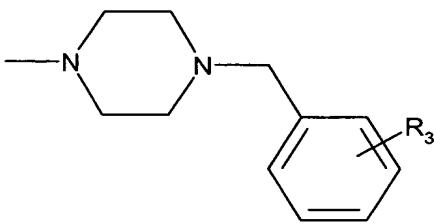
7 (currently amended): A compound according to claim 1 wherein L is CO and -Lp(D)n is of the formula:



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wherein;

m represents 0 or 1;

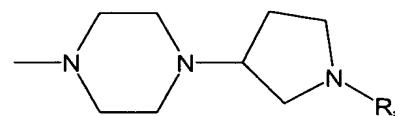
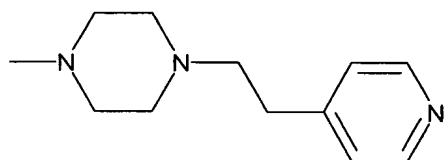
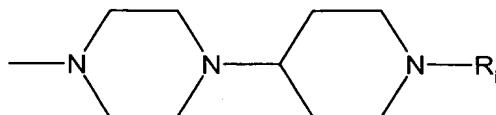
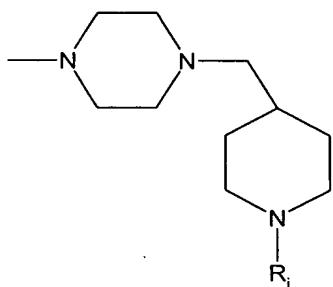
x⁰ represents CH or N; and

when R₃ is present as a substituent on an aromatic ring, it is selected from hydrogen, alkylsulphonyl, aminosulphonyl, alkylaminosulphonyl, alkylaminocarbonyl, amino, amido, alkoxy carbonyl, acetyl amino, chloro, fluoro, cyano, methoxy, ethoxy, nitro, hydroxy, alkylsulphonylamino, triazolyl and tetrazolyl; and

when R₃ is present as a substituent on a saturated ring, it is selected from hydrogen, hydroxy, amino, (1-3C) alkoxy, (1-3C) hydroxyalkyl, (1-3C) alkyl, carboxy, methoxycarbonyl and ethoxycarbonyl.

8 (original): A compound according to claim 7 wherein -Lp(D)n is of the formula:

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wherein R_i is hydrogen or (1-6C)alkyl.

9 (currently amended): A compound according to claim ~~any one~~ of claims 1 to 8 wherein R₂ is: - phenyl, thien-2-yl, naphthyl, indol-2-yl, indol-6-yl, benzo[b]furan-5-yl, benzo[b]thiophen-2-yl or benzimidazol-2-yl (each of which is optionally substituted as defined in claim 1).

(i) phenyl optionally being substituted in the 3 and/or 4 position by fluoro, chloro, bromo, iodo, nitro, difluoromethoxy, trifluoromethoxy, amino, cyano, trifluoromethyl, methylthio, vinyl, carboxy, acetoxy, MeSO₂⁻, hydroxy, methoxy, ethoxy, methyl, methoxycarbonyl, methylamino, ethylamino or amido, and optionally substituted at the 6 position by amino, hydroxy, fluoro, methoxycarbonyl, cyano or aminomethyl;

(ii) naphth-2-yl optionally substituted at the 6, position by hydroxy and optionally substituted at the 3 position by amino or hydroxy;

(iii) isoquinolin-7-yl, indol-5-yl, indol-6-yl, indazol-5-yl, indazol-6-yl, benzothiazol-6-yl or benzisoxazol-5-yl optionally substituted at the 3 position by chloro, bromo, amino, methyl or methoxy;

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- (iv) benzimidazol-5-yl or benzothiazol-6-yl optionally substituted at the 2 position by amino;
- (v) thien-2-yl or thien-3-yl optionally substituted at the 4 or 5 position by methylthio, methyl or acetyl;
- (vi) 3,4-methylenedioxypyphenyl, 2,3-dihydroindol-6-yl, 3,3-dichloro-2-oxo-indol-6-yl or 1-methyl-3-aminoindazol-5-yl;
- (vii) benzothiazol-2-yl, imidazo[1,2-a]pyrimidin-2-yl or tetrahydroimidazo[1,2-a]pyrimidin-2-yl;
- (viii) pyrazol-2-yl substituted at the 5 position by methyl;
- (ix) pyrid-2-yl optionally substituted at the 6 position by chloro;
- (x) pyrid-3-yl optionally substituted at the 4 position by chloro;
- (xi) benzofur-2-yl optionally substituted at the 3 position by chloro, methyl or methoxy, at the 5 or 6 position by methyl and at the 6 position by methoxy;
- (xii) indol-2-yl optionally substituted on the indole nitrogen atom by methyl and optionally substituted at the 5 or 6 position by fluoro, chloro, bromo, methyl or methoxy;
- (xiii) indol-6-yl substituted at the 5 position by chloro, fluoro or hydroxy and optionally substituted at the 3 position by chloro or methyl; or
- (xiv) benzo[b]thiophen-2-yl optionally substituted at the 3 position by fluoro, chloro or methyl, and optionally substituted at the 5 or 6 position by fluoro, chloro, methyl, hydroxy, or methoxy.

10 (currently amended): A compound according to claim ~~any one of claims 1 to 9 wherein optional substituents for R₂ are~~ is selected from:-

- (i) phenyl, 2-aminophenyl, 3-aminophenyl, 2-amino-3-fluorophenyl, 2-amino-4-fluorophenyl, 2-amino-4-chlorophenyl,

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2-amino-3-bromophenyl, 2-amino-3-nitrophenyl, 2-amino-4-nitrophenyl, 3,4-dimethoxy-5-aminophenyl, 2-amino-4-methylphenyl, 2-amino-3-methylphenyl, 2-amino-3-methoxyphenyl, 3,4-diaminophenyl, 3,5-diaminophenyl, 3-amino-4-fluorophenyl, 3-amino-4-chlorophenyl, 3-amino-4-bromophenyl, 3-amino-4-hydroxyphenyl, 3-amino-4-carboxymethylphenyl, 3-amino-4-methylphenyl, 3-amino-4-methoxyphenyl, 2-fluorophenyl, 4-fluoro-3-cyanophenyl, 3-chlorophenyl, 3-chloro-4-hydroxphenyl, 3-chloro-5-hydroxyphenyl, 4-chlorophenyl, 4-chloro-2-hydroxyphenyl, 4-chloro-3-hydroxyphenyl, 4-chloro-3-methylphenyl, 4-chloro-3-methoxyphenyl, 4-bromophenyl, 4-bromo-3-methylphenyl, 4-iodophenyl, 2-cyanophenyl, 3-cyanophenyl, 4-cyanophenyl, 3-cyano-5-aminophenyl, 2-hydroxphenyl, 2-hydroxy-4-methoxyphenyl, 3-hydroxphenyl, 3-hydroxy-4-methylphenyl, 2,4-dihydroxyphenyl, 3,4-dihydroxyphenyl, 3-hydroxy-4-methoxyphenyl, 4-difluoromethoxyphenyl, 4-trifluoromethoxphenyl, 4-trifluoromethylphenyl, 4-methylthiophenyl, 4-methoxycarbonylphenyl, 4-acetoxyphenyl, 4-methanesulfonylphenyl, 3-methylphenyl, 3-methyl-5-aminophenyl, 4-methylphenyl, 4-vinylphenyl, 4-methoxyphenyl, 4-ethoxyphenyl, 4-methoxy-3-chlorophenyl, 4-methoxy-3-methylphenyl, 3-methylaminophenyl, 4-methylaminophenyl, 4-ethylaminophenyl or 2-aminomethylphenyl;

(ii) naphth-2-yl, 3-aminonaphth-2-yl, 3-hydroxynaphth-2-yl or 6-hydroxynaphth-2-yl;

(iii) isoquinolin-7-yl, indol-5-yl, indol-6-yl, 3-chloroindol-6-yl, 3-bromoindol-6-yl, 3-methylindol-6-yl, 3-methoxyindol-6-yl, indazol-5-yl, 3-aminoindazol-5-yl, indazol-6-yl, benzothiazol-6-yl, 3-aminobenzisoxazol-5-yl;

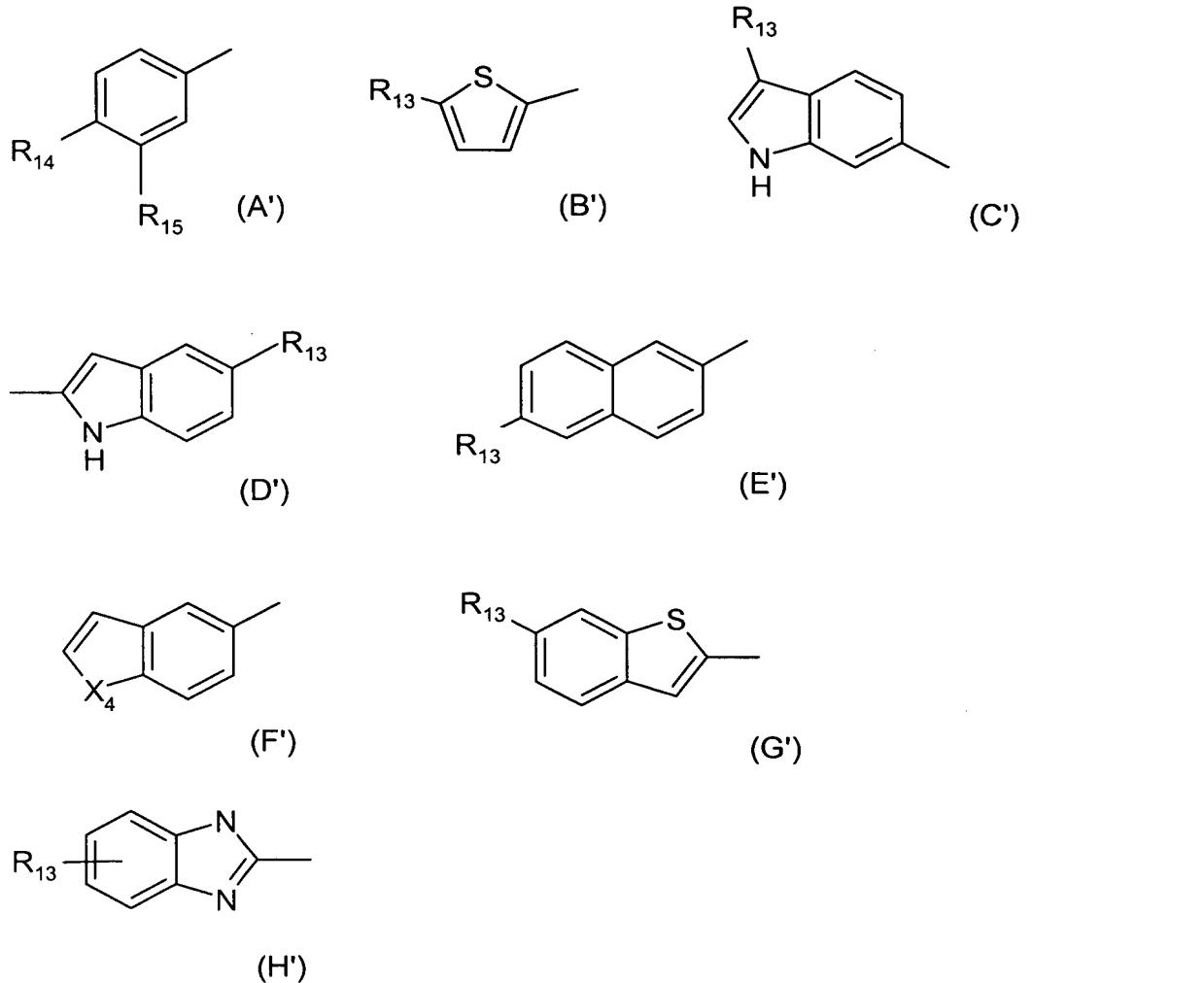
(iv) benzimidazol-5-yl, 2-aminobenzimidazol-5-yl, or benzothiazol-6-yl;

(v) thien-2-yl, 5-methylthien-2-yl, 5-methylthio-thien-2-yl, 5-acetylthien-2-yl or thien-3-yl;

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- (vi) 3,4-methylenedioxyphenyl, 2,3-dihydroindol-6-yl,
3,3-dichloro-2-oxo-indol-6-yl or 1-methyl-3-aminoindazol-5-yl;
- (vii) benzothiazol-2-yl, imidazo[1,2-a]pyrimidin-2-yl or
tetrahydroimidazo[1,2-a]pyrimidin-2-yl;
- (viii) 5-methylpyrazol-2-yl;
- (ix) 5-chloropyrid-2-yl;
- (x) pyrid-3-yl, 6-chloropyrid-3-yl;
- (xi) benzofur-2-yl, 5-chlorobenzofur-2-yl, 3-methylbenzofur-2-yl, 5-methylbenzofur-2-yl, 6-methoxybenzofur-2-yl;
- (xii) indol-2-yl, 5-fluoroindol-2-yl, 5-chloroindol-2-yl, 5-methylindol-2-yl, 5-methoxindol-2-yl, 6-methoxyindol-2-yl and 1-methyl-indol-2-yl;
- (xiii) 5-fluoroindol-6-yl; or
- (xiv) benzo[b]thiophen-2-yl, 5-chloro- benzo[b]thiophen-2-yl or 6-chlorobenzo[b]thiophen-2-yl.~~fluoro, chloro, bromo, iodo, nitro, thiol, difluoromethoxy, trifluoromethoxy, hydrazido, methylhydrazido, amino, cyano, trifluoromethyl, methylthio, vinyl, ethynyl, acetyl amino, carboxy, acetoxy, hydroxy, methyl, ethyl, amide (CONH₂), aminomethyl, methoxy and ethoxy.~~

11 (currently amended): A compound according to claim ~~any one~~ of claims 1 to 10 wherein R₂ is selected from one of the formula (A') to (H'):



wherein X₄ is O or S, R₁₃ is selected from hydrogen, fluoroo-, [except for (C')], chloro or methyl and R₁₄ is selected from hydrogen, methyl, ethyl, fluoro, chloro, and methoxy and R₁₅ is selected from hydrogen, methyl, fluoro, chloro and amino.

12 (currently amended): A compound according to claim ~~claims~~
~~1-10-11~~, wherein R₂ is 4-chlorophenyl, 4-methoxyphenyl, 3-amino-4-chlorophenyl, indol-2-yl, 5-chloroindol-2-yl, indol-6-yl, 3-chloroindol-6-yl or 3-methylindol-6-yl.

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13 (canceled) :

14 (canceled) :

15 (canceled) :

16 (canceled) :

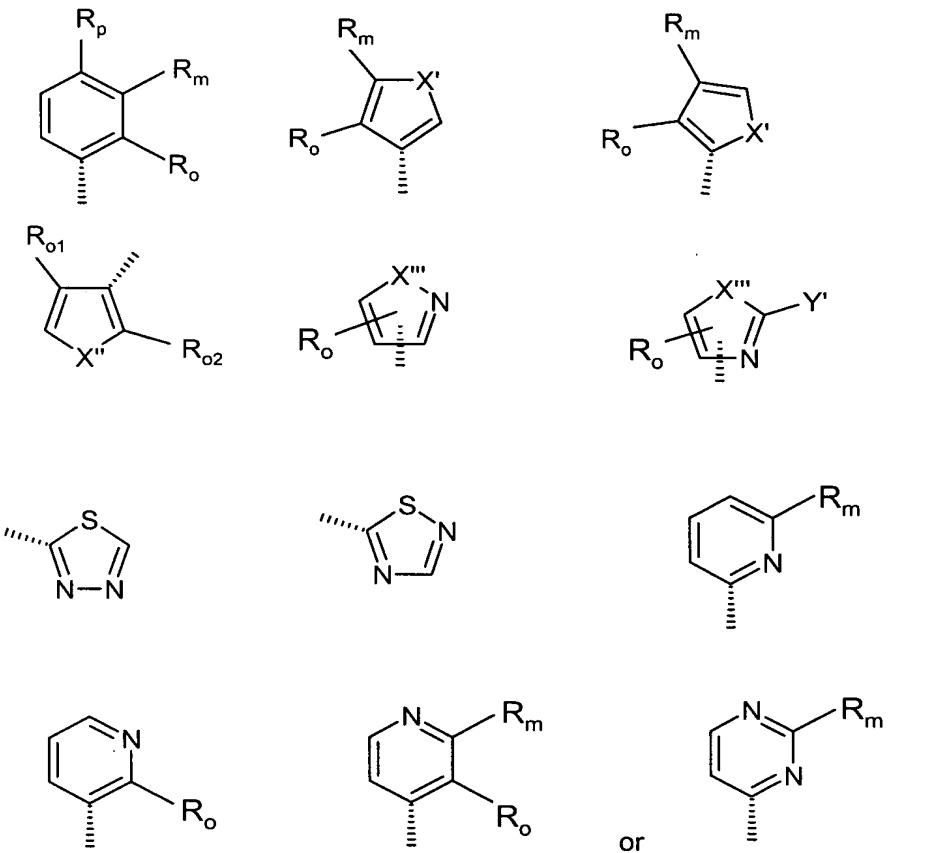
17 (canceled) :

18 (canceled) :

19 (currently amended) : A compound according to claim 1 ~~any one of claims 1 to 16~~ wherein R_{3a} is selected from hydrogen, hydroxyl, methoxy, ethoxy, methyl, ethyl, methylaminomethyl, dimethylaminomethyl, hydroxymethyl, carboxy, methoxymethyl, methoxycarbonyl, ethoxycarbonyl, methylaminocarbonyl, dimethylamino-carbonyl, aminomethyl, CONH₂, CH₂CONH₂, acetylarnino, methoxycarbonylamino, ethoxycarbonylamino, t-butoxycarbonylamino, amino, fluoro, chloro, bromo, cyano, nitro, thiol, methylthio, methylsulphonyl, ethylsulphonyl, methylsulphenyl, methylsulphonylamido, ethylsulphonylamido, methylaminosulphonyl, ethylaminosulphonyl, aminosulphonyl, trifluoromethoxy, trifluoromethyl, bromo, -OCH₂O- (which is bonded to two adjacent ring atoms in Cy) and -C(X³)N(R¹¹)R¹² (wherein X³ is O or S and R¹¹ and R¹² are independently selected from hydrogen, methyl or ethyl or together with the nitrogen atom to which they are attached form a pyrrolidin-1-yl, piperidin-1-yl or morpholino group).

20 (canceled) :

21 (currently amended) : A compound according to claim ~~any one of claims 1 to 14~~ wherein Cy is selected from:



wherein:

X' is selected from O, S and NMe;

X'' is selected from O and S;

X''' is selected from O, S, NH and NMe;

Y' is selected from hydrogen, amino and methyl;

R_o is selected from hydrogen, methyl, fluoro, chloro, trifluoromethyl, methoxy, methylthio, methylsulphinyl and methylsulphonyl;

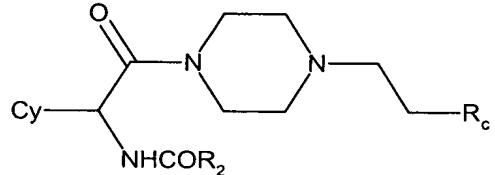
R_m is selected from hydrogen, methyl, fluoro, chloro, trifluoromethyl, methoxy, methylthio, methylsulphinyl, methylsulphonyl, carboxy, methoxycarbonyl and a group of the

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formula $-C(X^3)N(R^{11})R^{12}$ (wherein X^3 is O or S and R^{11} and R^{12} are independently selected from hydrogen, methyl or ethyl or together with the nitrogen atom to which they are attached form a pyrrolidin-1-yl, piperidin-1-yl or morpholino group); R_p is selected from hydrogen and fluoro; or R_O and R_m or R_m and R_p form an $-OCH_2O-$ group; or R_O and R_m together with the ring to which they are attached form a 5 or 6 membered aryl or heteroaryl ring (wherein the heteroaryl ring contains 1 or 2 heteroatoms selected from nitrogen, oxygen and sulfur); and one of R_{O1} and R_{O2} is hydrogen and the other is $R_{O\oplus}$.

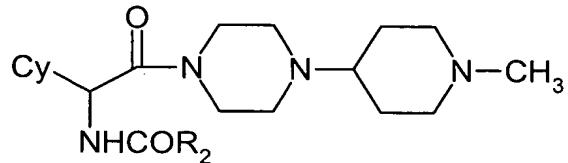
22 (currently amended): A compound according to claim ~~any one~~ of claims 1 to 14 wherein Cy is selected from phenyl, 2-chlorophenyl, 2-methoxyphenyl, 4-carbamoylphenyl, pyrid-2-yl, pyrid-4-yl, thien-2-yl, thien-3-yl, furan-2-yl, furan-3-yl, imidazol-2-yl, thiazol-2-yl, thiazol-4-yl, thiazol-5-yl and quinolin-4-yl.

23 (currently amended): A compound of the formula:



wherein Cy, R₂ and R_c are as defined in any one of claims 1 to 22, 9 to 12, 19 and 21 to 22.

24 (currently amended): A compound of the formula:



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wherein Cy and R₂ are as defined in any one of claims 1 to 22, 9 to 12, 19 and 21 to 22.

25 (canceled) :

26 (currently amended) : A compound as claimed in Claim 1, which is selected from:

1-(Indole-6-carbonyl-D-phenylglycinyl)-4-[2-(4-pyridinyl)-ethyl]piperazine;
1-(3-Chloroindole-6-carbonyl-D-phenylglycinyl)-4-[2-(4-pyridinyl)ethyl]piperazine;
1-(4-Methoxybenzoyl-D-phenylglycinyl)-4-(1-methylpiperidin-4-yl)piperazine;
~~1-(Indole-6-carbonyl-D-phenylglycinyl)-4-(1-methylpiperidin-4-yl)piperazine;~~
1-(4-Methoxybenzoyl-D-(2-chlorophenyl)glycinyl)-4-(1-methylpiperidin-4-yl)piperazine;
1-(Indole-6-carbonyl-D-(2-chlorophenyl)glycinyl)-4-(1-methylpiperidin-4-yl)piperazine; and
1-(4-Methoxybenzoyl-D-(2-trifluoromethylphenyl)glycinyl)-4-(1-methylpiperidin-4-yl)piperazine;
and physiologically-tolerable salts thereof.

27 (currently amended) : A pharmaceutical composition, which comprises a compound as claimed in claim ~~any one of claims 1 to 26~~ together with at least one pharmaceutically acceptable carrier or excipient.

28 (canceled) :

29 (canceled) :

30 (currently amended) : A method of treatment of a human or non-human animal body to combat a thrombotic disorder selected

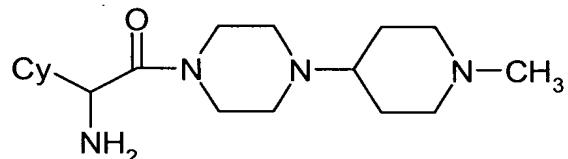
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from venous thrombosis, pulmonary embolism, arterial thrombosis, myocardial ischaemia, myocardial infarction and cerebral thrombosis, which comprises administering to said body an effective amount of a compound as claimed in claim 1.

31 (canceled) :

32 (canceled) :

33 (currently amended) : A compound of the formula



or a salt thereof in which Cy is as defined in any one of claims 1, 21 and 22.

34 (new) : A compound as claimed in any one of claims 1 to 13, 19 and 21 to 22, wherein the alpha atom in Y is carbon and has the conformation that would result from construction from a D- α -aminoacid NH₂-CH(Cy)-COOH where the NH₂ represents part of X-X.

35 (new) : A pharmaceutical composition, which comprises a compound as claimed in claim 34 together with at least one pharmaceutically acceptable carrier or excipient.

36 (new) : A method of treatment of a human or non-human animal body to combat a thrombotic disorder selected from venous thrombosis, pulmonary embolism, arterial thrombosis, myocardial ischaemia, myocardial infarction and cerebral thrombosis, which comprises administering to said body an effective amount of a compound as claimed in claim 34.

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37 (new): A method as claimed in claim 36 in which said human or non-human animal body is a human body.